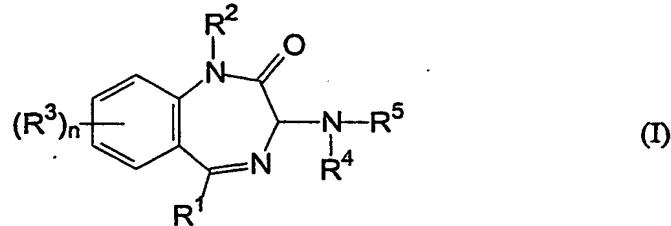


CLAIMS

1. Use of a benzodiazepine derivative of formula (I), or a pharmaceutically acceptable salt thereof, in the manufacture of a medicament for use in treating or preventing an RSV infection

5



wherein:

- R¹ represents C₁₋₆ alkyl, aryl or heteroaryl;
- R² represents hydrogen or C₁₋₆ alkyl;
- 10 - each R³ is the same or different and represents halogen, hydroxy, C₁₋₆ alkyl, C₁₋₆ alkoxy, C₁₋₆ alkylthio, C₁₋₆ haloalkyl, C₁₋₆ haloalkoxy, amino, mono(C₁₋₆ alkyl)amino, di(C₁₋₆ alkyl)amino, nitro, cyano, -CO₂R', -CONR'R'', -NH-CO-R', -S(O)R', -S(O)₂R', -NH-S(O)₂R', -S(O)NR'R'' or -S(O)₂NR'R'', wherein each R' and R'' is the same or different and represents hydrogen or C₁₋₆ alkyl;
- 15 - n is from 0 to 3;
- R⁴ represents hydrogen or C₁₋₆ alkyl;
- R⁵ represents C₁₋₆ alkyl, aryl, heteroaryl, carbocyclyl, heterocyclyl, aryl-(C₁₋₆ alkyl)-, heteroaryl-(C₁₋₆ alkyl)-, carbocyclyl-(C₁₋₆ alkyl)-, heterocyclyl-(C₁₋₆ alkyl)-, aryl-(C₁₋₆ hydroxyalkyl)-, heteroaryl-(C₁₋₆ hydroxyalkyl)-, carbocyclyl-(C₁₋₆ hydroxyalkyl)-, heterocyclyl-(C₁₋₆ hydroxyalkyl)-, aryl-C(O)-C(O)-, heteroaryl-C(O)-C(O)-, carbocyclyl-C(O)-C(O)-, heterocyclyl-C(O)-C(O)- or -XR⁶;
- 20 - X represents -CO-, -S(O)- or -S(O)₂-; and
- 25 - R⁶ represents C₁₋₆ alkyl, hydroxy, C₁₋₆ alkoxy, C₁₋₆ alkylthio, aryl, heteroaryl, carbocyclyl, heterocyclyl, aryl-(C₁₋₆ alkyl)-, heteroaryl-(C₁₋₆ alkyl)-, carbocyclyl-(C₁₋₆ alkyl)-, heterocyclyl-(C₁₋₆ alkyl)-, aryl-(C₁₋₆ alkyl)-

O-, heteroaryl-(C₁₋₆ alkyl)-O-, carbocyclyl-(C₁₋₆ alkyl)-O-, heterocyclyl-(C₁₋₆ alkyl)-O- or -NR'R'' wherein each R' and R'' is the same or different and represents hydrogen, C₁₋₆ alkyl, carbocyclyl, heterocyclyl, aryl, heteroaryl, aryl-(C₁₋₆ alkyl)-, heteroaryl-(C₁₋₆ alkyl)-, carbocyclyl-(C₁₋₆ alkyl)- or heterocyclyl-(C₁₋₆ alkyl)-.

2. Use according to claim 1 wherein:

- each R³ is the same or different and represents halogen, hydroxy, C₁₋₆ alkyl, C₁₋₆ alkoxy, C₁₋₆ alkylthio, C₁₋₆ haloalkyl, C₁₋₆ haloalkoxy, amino, mono(C₁₋₆ alkyl)amino, di(C₁₋₆ alkyl)amino, nitro, cyano, -CO₂R', -CONR'R'', -NH-CO-R', -S(O)R', -S(O)₂R', -NH-S(O)₂R' or -S(O)NR'R'', wherein each R' and R'' is the same or different and represents hydrogen or C₁₋₆ alkyl;
- R⁵ represents C₁₋₆ alkyl, aryl, heteroaryl, carbocyclyl, heterocyclyl, aryl-(C₁₋₆ alkyl)-, heteroaryl-(C₁₋₆ alkyl)-, carbocyclyl-(C₁₋₆ alkyl)-, heterocyclyl-(C₁₋₆ alkyl)- or -XR⁶;
- X represents -CO-, -S(O)- or -S(O)₂-; and
- R⁶ represents C₁₋₆ alkyl, hydroxy, C₁₋₆ alkoxy, C₁₋₆ alkylthio, aryl, heteroaryl, carbocyclyl, heterocyclyl, aryl-(C₁₋₆ alkyl)-, heteroaryl-(C₁₋₆ alkyl)-, carbocyclyl-(C₁₋₆ alkyl)-, heterocyclyl-(C₁₋₆ alkyl)- or -NR'R'' wherein each R' and R'' is the same or different and represents hydrogen, C₁₋₆ alkyl, carbocyclyl, heterocyclyl, aryl, heteroaryl, aryl-(C₁₋₆ alkyl)- or heteroaryl-(C₁₋₆ alkyl)-.

25. 3. Use according to either claim 1 or claim 2, wherein R¹ is C₁₋₂ alkyl or aryl.

4. Use according to any one of the preceding claims wherein R² is hydrogen.

30. 5. Use according to any one of the preceding claims wherein R³ is halogen, hydroxy, C₁₋₄ alkyl, C₁₋₄ alkoxy, C₁₋₄ alkylthio, C₁₋₄ haloalkyl, C₁₋₄ haloalkoxy, amino, mono(C₁₋₄ alkyl)amino or di(C₁₋₄ alkyl)amino.

6. Use according to claim 5, wherein R³ is fluorine, chlorine, bromine, C₁₋₂ alkyl, C₁₋₂ alkoxy, C₁₋₂ alkylthio, C₁₋₂ haloalkyl, C₁₋₂ haloalkoxy, amino, mono(C₁₋₂ alkyl)amino or di (C₁₋₂ alkyl)amino.
- 5 7. Use according to any one of the preceding claims wherein R⁴ is hydrogen or C₁₋₂ alkyl.
8. Use according to any one of the preceding claims wherein R⁵ is C₁₋₆ alkyl, aryl, heteroaryl, carbocyclyl, heterocyclyl, aryl-(C₁₋₄ alkyl)-, heteroaryl-(C₁₋₄ alkyl)-, carbocyclyl-(C₁₋₄ alkyl)-, heterocyclyl-(C₁₋₄ alkyl)-, aryl-C(O)-C(O)-, heteroaryl-C(O)-C(O)- or -XR⁶.
- 10 9. Use according to claim 8, wherein R⁵ is C₁₋₄ alkyl, aryl, heteroaryl, carbocyclyl, heterocyclyl, phenyl-(C₁₋₂ alkyl)-, heteroaryl-(C₁₋₂ alkyl)-, phenyl-C(O)-C(O)-, heteroaryl-C(O)-C(O)- or -XR⁶.
- 15 10. Use according to claim 9, wherein R⁵ is C₁₋₄ alkyl, phenyl, thienyl, furanyl, isoxazolyl, pyridyl, cyclopentyl, cyclohexyl, benzothienyl, dihydrobenzofuranyl, phenyl-CH₂-, furanyl-CH₂-, phenyl-C(O)-C(O)-, thienyl-C(O)-C(O)- or -XR⁶.
- 20 11. Use according to any one of the preceding claims wherein X is -CO- or -S(O)₂-.
- 25 12. Use according to any one of the preceding claims wherein, when R⁶ is a group -NR'R'' wherein each R' and R'' is the same or different and represents hydrogen, C₁₋₄ alkyl, aryl, carbocyclyl, heterocyclyl, aryl-(C₁₋₄ alkyl)- or heteroaryl-(C₁₋₄ alkyl)-.
- 30 13. Use according to claim 12, wherein when R⁶ is a group -NR'R'' each R' and R'' is the same or different and represents hydrogen, C₁₋₄ alkyl, phenyl, thienyl, cyclohexyl, cyclopentyl or phenyl-CH₂-.

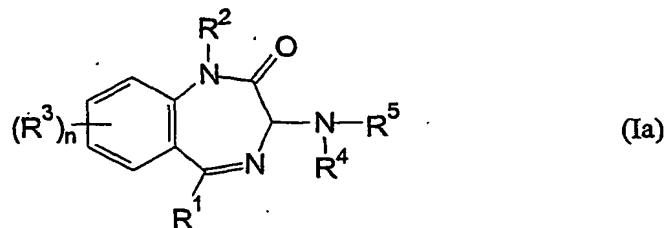
14. Use according to claim 13, wherein when R^6 is a group $-NR'R''$ and one of R' and R'' is hydrogen.

5 15. Use according to any one of the preceding claims wherein R^6 is C_{1-6} alkyl, hydroxy, C_{1-6} alkoxy, C_{1-6} alkylthio, aryl, heteroaryl, carbocyclyl, heterocyclyl, aryl- $(C_{1-4}$ alkyl)-, heteroaryl- $(C_{1-4}$ alkyl)-, carbocyclyl- $(C_{1-4}$ alkyl)-, heterocyclyl- $(C_{1-4}$ alkyl)-, aryl- $(C_{1-4}$ hydroxyalkyl)-, heteroaryl- $(C_{1-4}$ hydroxyalkyl)-, carbocyclyl- $(C_{1-4}$ hydroxyalkyl)-, heterocyclyl- $(C_{1-4}$ hydroxyalkyl)-, aryl- $(C_{1-4}$ alkyl)- O -, heteroaryl- $(C_{1-4}$ alkyl)- O -, carbocyclyl- $(C_{1-4}$ alkyl)- O -, heterocyclyl- $(C_{1-4}$ alkyl)- O - or $-NR'R''$.

10 16. Use according to claim 15, wherein R^6 is C_{1-6} alkyl, C_{1-6} alkoxy, C_{1-6} alkylthio, aryl, heteroaryl, carbocyclyl, heterocyclyl, phenyl- $(C_{1-2}$ alkyl)-, phenyl- $(C_{1-2}$ alkyl)- O -, heteroaryl- $(C_{1-2}$ alkyl)-, phenyl- $(C_{1-2}$ hydroxyalkyl)-, heteroaryl- $(C_{1-2}$ hydroxyalkyl)- or $-NR'R''$.

15 17. Use according to claim 16, wherein R^6 is C_{1-4} alkyl, C_{1-4} alkoxy, phenyl, naphthyl, dihydrobenzofuranyl, benzodioxinyl, 9H-fluoren-9-onyl, indolyl, thienyl, furanyl, oxazolyl, isoxazolyl, pyrazolyl, pyridyl, benzothienyl, benzofuranyl, cyclopentyl, cyclohexyl, piperazinyl, piperidinyl, morpholinyl, phenyl- $(C_{1-2}$ alkyl)-, phenyl- $CH_2-CH(OH)-$, phenyl- $CH(OH)-CH_2-$, phenyl- $(C_{1-2}$ alkyl)- O -, 1H-benzo[*d*]imidazol-2(3*H*)-onyl or $-NR'R''$.

20 18. Use according to any one of the preceding claims wherein the benzodiazepine derivative of formula (I) is a benzodiazepine derivative of formula (Ia):



wherein:

- R^1 is phenyl or methyl;
- R^3 is methyl or chlorine;
- 5 - n is 0 or 1;
- R^4 is hydrogen or methyl;
- R^5 is phenyl- CH_2 -, furanyl- CH_2 -, thieryl-C(O)-C(O)- or - XR^6 ;
- X is -CO- or -S(O)₂-; and
- R^6 is C_{1-4} alkyl, C_{1-4} alkoxy, phenyl, naphthyl, dihydrobenzofuranyl, benzodioxinyl, 9H-fluoren-9-onyl, indolyl, thieryl, furanyl, oxazolyl, 10 isoxazolyl, pyrazolyl, pyridyl, benzothienyl, benzofuranyl, cyclopentyl, cyclohexyl, piperazinyl, piperidinyl, morpholinyl, phenyl-(C_{1-2} alkyl)-, phenyl- CH_2 - $CH(OH)$ -, phenyl- $CH(OH)$ - CH_2 -, phenyl-(C_{1-2} alkyl)-O-, 1H-benzo[d]imidazol-2(3H)-onyl or - $NR'R''$ wherein each R' and R'' is the same 15 or different and represents hydrogen, C_{1-4} alkyl, phenyl, thieryl, cyclohexyl, cyclopentyl or phenyl-(CH_2)-,

the phenyl moiety in the group R^1 being unsubstituted or substituted by a single fluorine, chlorine, C_{1-2} alkyl, C_{1-2} alkoxy, C_{1-2} alkylthio, C_{1-2} haloalkyl or C_{1-2} haloalkoxy substituent;

20 the aryl moieties in the groups R^5 and R^6 being unsubstituted or substituted by 1,2 or 3 substituents selected from fluorine, chlorine, bromine, iodine, C_{1-4} alkyl, C_{2-4} acyl, hydroxy, C_{1-4} alkoxy, C_{1-4} alkylthio, C_{1-4} haloalkyl, C_{1-4} haloalkoxy, amino, mono(C_{1-4} alkyl)amino, di(C_{1-4} alkyl)amino, nitro, - CO_2R' , - $S(O)_2R'$ and - $S(O)_2NH_2$, wherein R' represents C_{1-2} alkyl;

25 the heteroaryl moieties in the groups R^5 and R^6 being unsubstituted or substituted by 1 or 2 substituents selected from fluorine, chlorine, bromine, C_{1-2} alkyl, C_{1-2} haloalkyl and di(C_{1-2} alkyl)amino; and

30 the heterocyclyl and carbocyclyl moieties in the R^6 group being unsubstituted or substituted by 1 or 2 substituents selected from fluorine, chlorine, bromine, C_{1-4} alkyl, C_{1-4} alkoxy, C_{1-4} haloalkyl and nitro.

19. Use according to any one of the preceding claims, wherein the medicament is for use in treating a patient who is a child under two years of age.
20. Use according to claim 19 wherein said child suffers from chronic lung disease.
21. Use according to any one of claims 1 to 18 wherein the medicament is for use in preventing RSV infection in an infant less than six years of age who was born after 32 weeks of gestation or less.
22. Use according to any one of the preceding claims, wherein the medicament is suitable for intranasal or intrabronchial administration.
23. Use according to any one of the preceding claims, wherein the medicament further comprises an anti-inflammatory compound or an anti-influenza compound.
24. Use according to claim 23 wherein the anti-inflammatory compound is budesonide or fluticasone.
25. Use according to claim 23 wherein the anti-inflammatory compound is a leukotriene antagonist, phosphodiesterase 4 inhibitor or TNF alpha inhibitor.
26. Use according to claim 23 wherein the anti-inflammatory compound is an interleukin 8 or interleukin 9 inhibitor.
27. Use according to any one of claims 1 to 22 wherein the medicament is coadministered with an anti-inflammatory compound, as defined in any one of claims 24 to 26, or an anti-influenza compound.
28. A method of treating a patient suffering from or susceptible to an RSV infection, which method comprises administering to said patient an effective

amount of a benzodiazepine derivative of formula (I), as defined in any one of claims 1 to 19, or a pharmaceutically acceptable salt thereof.

29. A method according to claim 28, wherein said patient is a patient as defined
5 in any one of claims 19 to 21.

30. A method according to claim 28 or 329, wherein the benzodiazepine derivative or salt thereof is administered intranasally or intrabronchially.

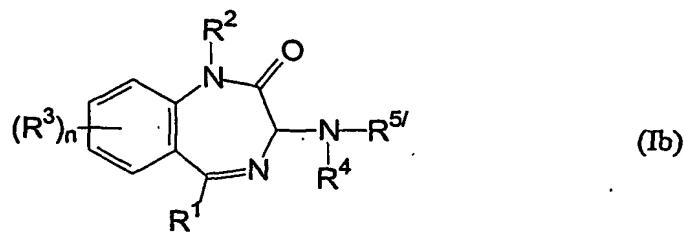
10 31. An inhaler or nebuliser containing a medicament which comprises
(a) a benzodiazepine derivative of formula (I), as defined in any one of claims 1 to 18, or a pharmaceutically acceptable salt thereof, and
(b) a pharmaceutically acceptable carrier or diluent.

15 32. A product comprising a compound of formula (I), or pharmaceutically acceptable salt thereof, as defined in any one of claims 1 to 18 and an anti-influenza compound, as defined in any one of claims 24 to 26, or an anti-influenza compound.

20 33. Use of a product according to claim 32 in the manufacture of a medicament for use in the treatment of concomitant RSV and influenza infections.

34. Use of a compound of formula (I), or pharmaceutically acceptable salt thereof, as defined in any one of claims 1 to 18 in the manufacture of a
25 medicament for use in the treatment of human metapneumovirus, measles, parainfluenza viruses, mumps, yellow fever virus (B5 strain), Dengue 2 virus or West Nile virus.

35. A benzodiazepine derivative of formula (Ib), or a pharmaceutically acceptable salt thereof
30



wherein:

- R¹ represents C₁₋₆ alkyl, aryl or heteroaryl;
- R² represents hydrogen, C₁₋₆ alkyl;
- 5 - each R³ is the same or different and represents halogen, hydroxy, C₁₋₆ alkyl, C₁₋₆ alkoxy, C₁₋₆ alkylthio, C₁₋₆ haloalkyl, C₁₋₆ haloalkoxy, amino, mono(C₁₋₆ alkyl)amino, di(C₁₋₆ alkyl)amino, nitro, cyano, -CO₂R', -CONR'R'', -NH-CO-R', -S(O)R', -S(O)₂R', -NH-S(O)₂R', -S(O)NR'R'' or -S(O)₂NR'R'', wherein each R' and R'' is the same or different and represents hydrogen or C₁₋₆ alkyl;
- 10 - n is from 0 to 3;
- R⁴ represents hydrogen or C₁₋₆ alkyl;
- R⁵ represents C₃₋₆ alkyl, aryl, heteroaryl, carbocyclyl, heterocyclyl, aryl-(C₁₋₆ alkyl)-, heteroaryl-(C₁₋₆ alkyl)-, carbocyclyl-(C₁₋₆ alkyl)-, heterocyclyl-(C₁₋₆ alkyl)-, aryl-C(O)-C(O)-, heteroaryl-C(O)-C(O)-, carbocyclyl-C(O)-C(O)-, heterocyclyl-C(O)-C(O)- or -X', provided that when R⁵ is heteroaryl it is not 2-quinaldyl or 6-chloro-pyrazinyl, when R⁵ is heteroaryl-(C₁₋₆ alkyl)- it is not 2-indolylmethyl, 2-(3-indolyl)ethyl or 2-furanylmethyl, when R⁵ is aryl it is not unsubstituted phenyl and when R⁵ is aryl-(C₁₋₆ alkyl)- it is not unsubstituted phenyl-(C₁₋₂ alkyl)- or 4-chlorophenyl-(C₂₋₃ alkyl)-;
- 15 - X' represents -CO-R⁶', -S(O)-R⁶'' or -S(O)₂-R⁶''';
- R⁶' represents C₁ alkyl, hydroxy, C₁₋₆ alkoxy, C₁₋₆ alkylthio, aryl, heteroaryl, carbocyclyl, heterocyclyl, aryl-(C₁₋₆ alkyl)-, heteroaryl-(C₁₋₆ alkyl)-, carbocyclyl-(C₁₋₆ alkyl)-, heterocyclyl-(C₁₋₆ alkyl)-, aryl-(C₁₋₆ alkyl)-O-, heteroaryl-(C₁₋₆ alkyl)-O-, carbocyclyl-(C₁₋₆ alkyl)-O-, heterocyclyl-(C₁₋₆ alkyl)-O- or -NR'R'' wherein each R' and R'' is the same or different and
- 20
- 25

represents hydrogen, C₁₋₆ alkyl, carbocyclyl, heterocyclyl, aryl, heteroaryl, aryl-(C₁₋₆ alkyl)-, heteroaryl-(C₁₋₆ alkyl)-, carbocyclyl-(C₁₋₆ alkyl)- or heterocyclyl-(C₁₋₆ alkyl)-, provided that (a) when R^{6/} is aryl it is not unsubstituted naphthyl, unsubstituted phenyl, mono-halophenyl, 5. 4-methylphenyl, 4-methoxyphenyl, 4-hydroxyphenyl, 4-trifluoromethylphenyl, 4-nitrophenyl, 4-cyanophenyl, 4-n-propylphenyl, 4-t-butylphenyl, 4-n-pentylphenyl, 4-dimethylaminophenyl, 4-methylthiophenyl, 3-trifluoromethylthiophenyl, 3,4-dimethoxyphenyl, 3,4-dichlorophenyl, 3,5-dichlorophenyl, 2,3,4,5,6-pentafluorophenyl, 4-chloro-2-aminophenyl or 4-10. 1,1-dimethylethylphenyl, (b) when R^{6/} is heteroaryl it is not 2-pyrrolyl, 2-pyrazinyl, 2-quinaldyl, 2-quinoxalinyl, 1-methylindolony, 2-methyl-indolyl, 2-benzofuranyl, 2-benzothienyl, 3-thienyl, 3-indolyl, unsubstituted 2-indolyl, 5-fluoroindol-2-yl, 5-chloroindol-2-yl, 5-bromoindol-2-yl, 5-hydroxyindol-2-yl or 5-methoxyindol-2-yl, (c) when R^{6/} is aryl-(C₁₋₆ alkyl)- it is not 4-15. thianaphthene-(CH₂)-, unsubstituted phenyl-(CH₂)-, 4-trifluoromethylphenyl-(CH₂)-, unsubstituted phenyl-(CH₂)₃-, monotrifluoromethylphenyl-(CH₂)₂-, 3-methoxyphenyl-(CH₂)₂-, 4-chloro-2-aminophenyl-(CH₂)₂-, 2,4-20. dichlorophenyl-(CH₂)₂-, monochlorophenyl-(CH₂)₂-, 2,4-trifluoromethylphenyl-(CH₂)₂-, 4-cyanophenyl-(CH₂)₂- or 3-cyanophenyl-(CH₂)₂-, (d) when R^{6/} is heteroaryl-(C₁₋₆ alkyl)- it is not indolyl-(CH₂)_x-, wherein x is 1, 2, 3, unsubstituted furanyl-(CH₂)₂-, unsubstituted thienyl-(CH₂)₃- (e) when R^{6/} is carbocyclyl it is not cyclohexyl, (f) when R^{6/} is carbocyclyl-(C₁₋₆ alkyl)- it is not unsubstituted cyclohexyl-(CH₂)₁₋₃-, (g) when R^{6/} is heterocyclyl it is not N-pyrrolidinyl or 2-dihydrobenzofuranyl, (h) when R^{6/} is aryl-(C₁₋₆ alkyl)-O-25. it is not unsubstituted phenyl-(CH₂)-O-, and (i) when R^{7/} is hydrogen, R^{8/} is not unsubstituted phenyl, 4-halophenyl, 3-halophenyl, methoxyphenyl, nitrophenyl, 2-chlorophenyl, 4-methylphenyl, dichlorophenyl, 3,5-dimethylphenyl, 3-methylphenyl, 3-cyanophenyl, 3-aminophenyl, 3-aminocarbonylphenyl, 3-benzoic acid, 3-benzoic acid ethyl ester, 6-amino-3-pyridyl, 5-(2-chloro)pyridyl, 5-(2-methoxy)pyridyl, 5-indanyl, unsubstituted cyclohexyl, 1,1-dimethylethyl, unsubstituted phenyl-CH₂-, unsubstituted naphthyl or benzotriazol-3-yl and when R^{7/} is methyl, R^{8/} is not 30.

cyclopropylbenzene;

- $R^{6//}$ represents C_{1-6} alkyl, hydroxy, C_{1-6} alkoxy, C_{1-6} alkylthio, aryl, heteroaryl, carbocyclyl, heterocyclyl, aryl-(C_{1-6} alkyl)-, heteroaryl-(C_{1-6} alkyl)-, carbocyclyl-(C_{1-6} alkyl)-, heterocyclyl-(C_{1-6} alkyl)-, aryl-(C_{1-6} alkyl)-O-, heteroaryl-(C_{1-6} alkyl)-O-, carbocyclyl-(C_{1-6} alkyl)-O-, heterocyclyl-(C_{1-6} alkyl)-O- or - $NR'R^{6//}$ wherein each R' and $R^{6//}$ is the same or different and represents hydrogen, C_{1-6} alkyl, carbocyclyl, heterocyclyl, aryl, heteroaryl, aryl-(C_{1-6} alkyl)-, heteroaryl-(C_{1-6} alkyl)-, carbocyclyl-(C_{1-6} alkyl)- or heterocyclyl-(C_{1-6} alkyl)-; and
- $R^{6//}$ represents C_{1-6} alkyl, hydroxy, C_{1-6} alkoxy, C_{1-6} alkylthio, aryl, heteroaryl, carbocyclyl, heterocyclyl, aryl-(C_{1-6} alkyl)-, heteroaryl-(C_{1-6} alkyl)-, carbocyclyl-(C_{1-6} alkyl)-, heterocyclyl-(C_{1-6} alkyl)-, aryl-(C_{1-6} alkyl)-O-, heteroaryl-(C_{1-6} alkyl)-O-, carbocyclyl-(C_{1-6} alkyl)-O-, heterocyclyl-(C_{1-6} alkyl)-O- or - $NR'R^{6//}$ wherein each R' and $R^{6//}$ is the same or different and represents hydrogen, C_{1-6} alkyl, carbocyclyl, heterocyclyl, aryl, heteroaryl, aryl-(C_{1-6} alkyl)-, heteroaryl-(C_{1-6} alkyl)-, carbocyclyl-(C_{1-6} alkyl)- or heterocyclyl-(C_{1-6} alkyl)-, provided that when $R^{6//}$ is aryl it is not 4-methylphenyl, provided that the compound of formula (Ib) is not N-(2-Oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-acetamide.

20

36. A benzodiazepine derivative according to claim 35 wherein:

- $R^{5/}$ is C_{3-6} alkyl, C_{3-6} cycloalkyl, heterocyclyl, C_{3-6} cycloalkyl-(C_{1-6} alkyl), aryl-C(O)-C(O)-, heteroaryl-C(O)-C(O)-, carbocyclyl-C(O)-C(O)-, heterocyclyl-C(O)-C(O)- or - X' ;
- X' is -CO- $R^{6/}$, -S(O)- $R^{6//}$ or -S(O)₂- $R^{6//}$;
- $R^{6/}$ is C_1 alkyl, hydroxy, C_{1-6} alkoxy, C_{1-6} alkylthio, heterocyclyl-(C_{1-6} alkyl)-, heteroaryl-(C_{1-6} alkyl)-O-, carbocyclyl-(C_{1-6} alkyl)-O-, heterocyclyl-(C_{1-6} alkyl)-O- or - $NR'R^{6//}$ wherein each R' and $R^{6//}$ is the same or different and represents hydrogen, C_{1-6} alkyl, C_{3-6} cycloalkyl, heterocyclyl, carbocyclyl-(C_{1-6} alkyl)- or heterocyclyl-(C_{1-6} alkyl)-;
- $R^{6//}$ represents C_{1-6} alkyl, hydroxy, C_{1-6} alkoxy, C_{1-6} alkylthio, aryl, heteroaryl, carbocyclyl, heterocyclyl, aryl-(C_{1-6} alkyl)-, heteroaryl-(C_{1-6}

alkyl)-, carbocyclyl-(C₁₋₆ alkyl)-, heterocyclyl-(C₁₋₆ alkyl)-, aryl-(C₁₋₆ alkyl)-O-, heteroaryl-(C₁₋₆ alkyl)-O-, carbocyclyl-(C₁₋₆ alkyl)-O-, heterocyclyl-(C₁₋₆ alkyl)-O- or -NR'R'' wherein each R' and R'' is the same or different and represents hydrogen, C₁₋₃ alkyl, heterocyclyl, heteroaryl, heteroaryl-(C₁₋₆ alkyl)-, carbocyclyl-(C₁₋₆ alkyl)- or heterocyclyl-(C₁₋₆ alkyl)-; and

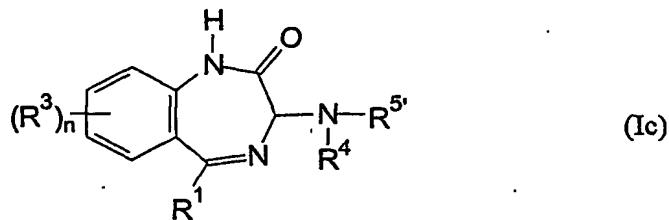
5 R^{6//} is C₁₋₆ alkyl, hydroxy, C₁₋₆ alkoxy, C₁₋₆ alkylthio, C₃₋₆ cycloalkyl, heterocyclyl, C₃₋₆ cycloalkyl-(C₁₋₆ alkyl)-, heterocyclyl-(C₁₋₆ alkyl)-, aryl-(C₁₋₆ alkyl)-O-, heteroaryl-(C₁₋₆ alkyl)-O-, carbocyclyl-(C₁₋₆ alkyl)-O-, heterocyclyl-(C₁₋₆ alkyl)-O- or -NR'R'' wherein each R' and R'' is the same or

10 different and represents hydrogen, C₁₋₆ alkyl, carbocyclyl, heterocyclyl, aryl, heteroaryl, aryl-(C₁₋₆ alkyl)-, heteroaryl-(C₁₋₆ alkyl), carbocyclyl-(C₁₋₆ alkyl)- or heterocyclyl-(C₁₋₆ alkyl)-.

37. A benzodiazepine derivative according to claim 35 or claim 36 wherein R² is

15 hydrogen.

38. A benzodiazepine derivative of formula (Ic), or a pharmaceutically acceptable salt thereof,



wherein:

25

- R¹ is phenyl or methyl;
- R³ is methyl or chlorine;
- n is 0 or 1;
- R⁴ is hydrogen or methyl;
- R^{5'} is phenyl-CH₂- thieryl-C(O)-C(O)- or -X';
- X' is -CO-R^{6'}, -CONR'R'', -S(O)₂R^{6'''} or -S(O)₂-NR₂ and

- R^{6i} is C_1 alkyl, C_{1-4} alkoxy, benzodioxinyl, 9H-fluoren-9-onyl, furanyl, oxazolyl, isoxazolyl, pyrazolyl, pyridyl, cyclopentyl, piperazinyl, piperidinyl, morpholinyl, phenyl- CH_2 - $CH(OH)$ -, phenyl- $CH(OH)$ - CH_2 -, phenyl-(C_2 alkyl)-O- or 1*H*-benzo[*d*]imidazol-2(3*H*)-only;
- 5 - R^{6m} is C_{1-4} alkyl, C_{1-4} alkoxy, phenyl, naphthyl, dihydrobenzofuranyl, benzodioxinyl, 9H-fluoren-9-onyl, indolyl, thienyl, furanyl, oxazolyl, isoxazolyl, pyrazolyl, pyridyl, benzothienyl, benzofuranyl, cyclopentyl, cyclohexyl, piperazinyl, piperidinyl, morpholinyl, phenyl-(C_{1-2} alkyl)-, phenyl- CH_2 - $CH(OH)$ -, phenyl- $CH(OH)$ - CH_2 -, phenyl-(C_{1-2} alkyl)-O- or 1*H*-benzo[*d*]imidazol-2(3*H*)-only;
- 10 - each R' and R'' is the same or different and represents hydrogen, C_{1-4} alkyl, phenyl, thienyl, cyclohexyl, cyclopentyl or phenyl-(CH_2)-; and
- each R_1 and R_2 is the same or different and represents hydrogen, C_{1-4} alkyl, phenyl, thienyl, cyclohexyl, cyclopentyl or phenyl-(CH_2)-, wherein:
- 15 the phenyl moiety in the group R^1 being unsubstituted or substituted by a single fluorine, chlorine, C_{1-2} alkyl, C_{1-2} alkoxy, C_{1-2} alkylthio, C_{1-2} haloalkyl or C_{1-2} haloalkoxy substituent;
- the aryl moieties in the groups R^{5i} , R^{6i} and R^{6m} being unsubstituted or substituted by 1,2 or 3 substituents selected from fluorine, chlorine, bromine, iodine, C_{1-4} alkyl, C_{2-4} acyl, hydroxy, C_{1-4} alkoxy, C_{1-4} alkylthio, C_{1-6} haloalkyl, C_{1-4} haloalkoxy, amino, mono(C_{1-4} alkyl)amino, di(C_{1-4} alkyl)amino, nitro, $-CO_2R'$, $-S(O)_2R'$ and $-S(O)_2NH_2$, wherein R' represents C_{1-2} alkyl;
- 20 the heteroaryl moieties in the groups R^{5i} , R^{6i} and R^{6m} being unsubstituted or substituted by 1 or 2 substituents selected from fluorine, chlorine, bromine, C_{1-2} alkyl, C_{1-2} haloalkyl and di(C_{1-2} alkyl)amino;
- the heterocyclyl and carbocyclyl moieties in the R^{6m} group being unsubstituted or substituted by 1 or 2 substituents selected from fluorine, chlorine, bromine, C_{1-4} alkyl, C_{1-4} alkoxy, C_{1-4} haloalkyl and nitro;
- 25 the aryl, heteroaryl and carbocyclyl moieties in the R' and R'' being unsubstituted or substituted by one or two substituents selected from fluorine, chlorine, bromine, C_{1-2} alkyl, C_{1-2} alkoxy, C_{1-2} alkylthio, C_{1-2} haloalkyl and

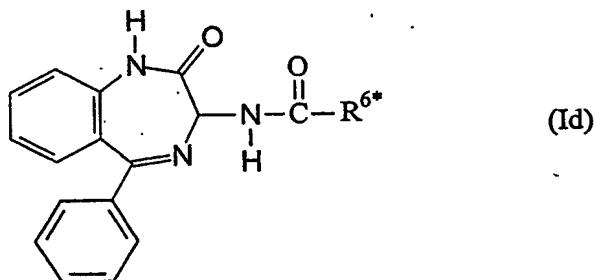
nitro; and

the aryl, heteroaryl and carbocyclyl moieties in the R₁ and R₂ being unsubstituted or substituted by one or two substituents selected from fluorine, chlorine, bromine, C₁₋₂ alkyl, C₁₋₂ alkoxy, C₁₋₂ alkylthio, C₁₋₂ haloalkyl and nitro,

5

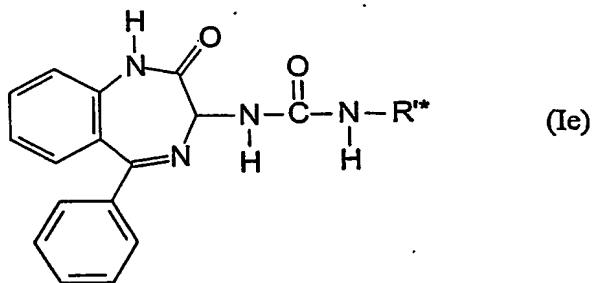
provided that the compound of formula (Ic) is not N-(2-Oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-acetamide.

10 39. A benzodiazepine derivative of formula (Id), or pharmaceutically acceptable salts thereof



15 wherein R^{6*} is an aryl group which is unsubstituted or substituted by 1, 2 or 3 substituents selected from halogen, C₁₋₆ alkyl, C₂₋₇ acyl, hydroxy, C₁₋₆ alkoxy, C₁₋₆ alkylthio, C₁₋₆ haloalkyl, C₁₋₆ haloalkoxy, nitro, cyano, carbamoyl, mono(C₁₋₆ alkyl)carbamoyl, di(C₁₋₆ alkyl)carbamoyl, amino, mono(C₁₋₆ alkyl)amino, di(C₁₋₆ alkyl)amino, -CO₂R', -CONR'R'', -S(O)R', -S(O)₂R', -S(O)NR'R'', -S(O)₂NR'R'' -NH-S(O)₂R' or -NH-CO-R', wherein each R' and R'' is the same or different and represents hydrogen or C₁₋₆ alkyl, provided that R^{6*} is not a 4-chlorophenyl group.

20 40. A benzodiazepine derivative of formula (Ie) or a pharmaceutically acceptable salts thereof



wherein R¹* is an aryl group which is unsubstituted or substituted by 1 or 2 substituents selected from fluorine, chlorine, bromine, C₁₋₄ alkyl, C₁₋₄ alkoxy, C₁₋₄ alkylthio, C₁₋₄ haloalkyl, C₁₋₄ haloalkoxy and nitro.

41. 1,1-Diethyl-3-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-urea
10 N-(2-Oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-propionamide
N-(2-Oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-butyramide
N-(2-Oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-isobutyramide
15 2,2-Dimethyl-N-(2-Oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-propionamide
Cyclopentanecarboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide
Cyclohexanecarboxylic acid 2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide.
20 Piperidine-1-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide
Morpholine-4-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide
4-Methyl-piperazine-1-carboxylic acid -(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide
25 Benzo[b]thiophene-3-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

Isoxazole-5-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

5 Benzo[b]thiophene-2-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

10 N-(2-Oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-methanesulfonamide

15 Propane-1-sulfonic acid-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

20 Butane-1-sulfonic acid-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

25 N-(7-Chloro-2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-isobutyramide

30 N-(2-Oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-isonicotinamide

35 N-(2-Oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-nicotinamide

40 N-(7-Chloro-2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-acetamide

45 (S)-2-Methoxy-4-nitro-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-benzamide

50 (S)-1-(2-Fluoro-phenyl)-3-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-urea

55 2-Chloro-4-methanesulfonyl-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-benzamide

60 1-(4-Nitro-phenyl)-3-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-urea

65 4-Methanesulfonyl-2-methoxy-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-benzamide

70 2-Methoxy-4-methylsulfanyl-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-benzamide

75 4-Methanesulfonyl-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-benzamide

N-(2-Oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)terephthalamic acid methyl ester

5-Acetyl-2-ethoxy-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-benzamide

5 3-Methoxy-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-terephthalamic acid methyl ester

2-Methylsulfanyl-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-benzamide

10 4-Amino-5-chloro-2-methoxy-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-benzamide

4-Methanesulfonyl-2-methoxy-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-benzamide

(S)-2,4,5-Trifluoro-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-benzamide

15 (S)-5-Acetyl-2-ethoxy-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-benzamide

2-Methoxy-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-5-sulfamoyl-benzamide

1-t-Butyl-3-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-urea

20 1-Cyclohexyl-3-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-urea

1-Ethyl-3-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-urea

1-Butyl-3-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-urea

25 4,5-Dimethyl-furan-2-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)amide

Piperidine-1-carboxylic acid (7-chloro-2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

N-[5-(3-Chloro-phenyl)-2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl]acetamide

30 N-[5-(3-Chloro-phenyl)-2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl]-isobutyramide

Cyclohexanecarboxylic acid [5-(3chloro-phenyl)-2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl]-amide

Piperidine-1-carboxylic acid [5-(3-chloro-phenyl)-2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl]-amide

5 N-[5-(3-Chloro-phenyl)-2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl]isonicotinamide

N-[5-(3-Methoxy-phenyl)-2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl]-isobutyramide

Cyclohexanecarboxylic acid [5-(3-methoxy-phenyl)-2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl]-amide

10 Piperidine-1-carboxylic acid [5-(3-methoxy-phenyl)-2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl]-amide

Piperidine-4-carboxylic acid [5-(3-methoxy-phenyl)-2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl]-amide

15 Cyclohexanecarboxylic acid (8-chloro-2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

6-Morpholin-4-yl-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-nicotinamide

Pyridine-2-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

20 6-Fluoro-4H-benzo[1,3]dioxine-8-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

1H-Pyrazole-4-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

25 6-Dimethylamino-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-nicotinamide

2-Ethoxy-naphthalene-1-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

9-Oxo-9H-fluorene-1-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

30 2-Oxo-2,3-dihydro-benzoimidazole-1-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

(2-Oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)carbamic acid
tert-butyl ester

(S)-6-Fluoro-4H-benzo[1,3]dioxine-8-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

5 (S)-4,5-Dibromo-furan-2-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

(S)-3-Methoxy-naphthalene-2-carboxylic acid (2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-amide

10 (2-Oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-carbamic acid methyl ester

(2-Oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-carbamic acid ethyl ester

(2-Oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-carbamic acid isobutyl ester

15 2-Oxo-N-(2-oxo-5-phenyl-2,3-dihydro-1H-benzo[e][1,4]diazepin-3-yl)-2-thiophene-2-yl-acetamide,
or a pharmaceutically acceptable salt thereof.

42. A benzodiazepine derivative according to any one of claims 35 to 41 for use
20 in a method of treating the human or animal body.

43. A pharmaceutical composition comprising a benzodiazepine derivative
according to any one of claims 35 to 41, or a pharmaceutically acceptable salt
thereof, and a pharmaceutically acceptable diluant or carrier.

25 44. A composition according to claim 43 comprising an optically active isomer of
a benzodiazepine derivative according to any one of claims 35 to 41.

45. A composition accoding to claim 43 or 44 which is in the form of a tablet,
30 troche, lozenge, aqueous or oily suspension, dispersible powders or granules.